## DT09 Rec'd PCT/PT0 0 9 SEP 2004

1/59

#### SEQUENCE LISTING

- <110> MASAI, Hisao

  TAMAI Katsuyuki
- <120> Cdc7-ASK KINASE COMPLEX, SUBSTRATES OF THE KINASE COMPLEX, SPECIFIC ANTIBODIES TO THE SUBSTRATES, AND SCREENING METHODS USING THE SAME TO SCREEN FOR COMPOUNDS COMPRISING Cdc7-ASK KINASE INHIBITORY ABILITY

<130> M3-A0201P

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<150> JP 2002-067702

<151> 2002-03-12

<160> 21

<170> PatentIn Ver. 2.1

<210> 1

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<212> PRT

<213> Homo sapiens

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tcc atc tat ggt cat gaa gac atc aag aga ggc cct gct ctg gcc ctg 1494  Ser Ile Tyr Gly His Glu Asp Ile Lys Arg Gly Pro Ala Leu Ala Leu 475 480 485  ttc gga ggg gag ccc aaa aac cca ggt ggc aag cac aag gta cgt ggt 1542  Phe Gly Gly Glu Pro Lys Asn Pro Gly Gly Lys His Lys Val Arg Gly 490 495 500  gat atc aac gtg ctc ttg tgc gga gac cct ggc aca gcg aag tcg cag 1590  Asp Ile Asn Val Leu Leu Cys Gly Asp Pro Gly Thr Ala Lys Ser Gln 505 510 515 520  ttt ctc aag tat att gag aaa gtg tcc agc cga gcc atc ttc acc act 1638  Phe Leu Lys Tyr Ile Glu Lys Val Ser Ser Arg Ala Ile Phe Thr Thr	Ser	Lys	Asp	Gln	Gln.	Ile	Gly	Glu	Lys	Ile	Phe	Ala	Ser	Ile	Ala	Pro	
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890

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		35					40					45	)		
Gly	Thr	Glu	Gly	Pro	Leu	Glu	Glu	Glu	Glu	Asp	Gly	Glu	Glu	Leu	Ile
•	50					55					60				
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65					70		•			<b>7</b> 5	٠				80
Tyr	Glu	Ala	Glu	G1y	Leu	Ala	Leu	Asp	Asp	Glu	Asp	Val	Glu	Glu	Leu
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Thr	Ala	Ser	Arg	Arg	Glu	Ala	Ala	Asp	Gly	Pro	Cys	G1y	Thr	Val	Thr
			100					105					110		
Gly	Ser	Trp	Pro	Gly	Leu	Gly	Ala	Cys	Ala	Val	Gly	Ser	Cys	Met	Thr
		115		•			120				٠	125			
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145					150					155					160
Glu	Asn	Leu	Glu	Asp	Leu	Lys	Gly	His	Ser	Val	Arg	Glu	Trp	Val	Ser
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Met	Ala	G1.y	Pro	Arg	Leu	Glu	Ile	His	His	Arg	Phe	Lys	Asn	Phe	Leu
			180	•				185				·.	190		
Arg	Thr	His	: Val	Asp	Ser	His	Gly	His	Asn	Val	Phe	Lys	Glu	Arg	Ile

		195					200					205			
Ser	Asp	Met	Cys	Lys.	Glu	Asn	Arg	Glu	Ser	Leu	Val	Val	Asn	Tyr	Glu
	210					215					220				
Asp	Leu	Ala	Ala	Arg	Glu	His	Val	Leu	Ala	Tyr	Phe	Leu	Pro	Glu	Ala
225					230					235				•	240
Pro	Ala	Glu	Leu	Leu	Gln	Ile	Phe	Asp	Glu	Ala	Ala	Leu	Glu	Val	Val
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Leu	Ala	Met	Tyr	Pro	Lys	Tyr	Asp	Arg	Ile	Thr	Asn	His	Ile	His	Val
			260					265					270		
Arg	Ile	Ser	His	Leu	Pro	Leu	Val	Glu	Glu	Leu	Arg	Ser	Leu	Arg	Gln
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Leu	His	Leu	Asn	Gln	Leu	Ile	Arg	Thr	Ser	Gly	Val	Val	Thr	Ser	Cys
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Thr	Gly	Val	Leu	Pro	Gln	Leu	Ser	Met	Val	Lys	Tyr	Asn	Cys	Asn	Lys
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Cys	Asn	Phe	Val	Leu	Gly	Pro	Phe	Cys	G1n	Ser	Gln	Asn	Gln	Glu	Val
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Lys	Pro	Gly	Ser	Cys	Pro	Glu	Cys	Gln	Ser	Ala	Gly	Pro	Phe	Glu	Val
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Asn	Met	Glu	Glu	Thr	Ile	Tyr	Gln	Asn	Tyr	Gln	Arg	Ile	Arg	Ile	Gln
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Glu	Ser	Pro	Gly	Lys	Val	Ala	Ala	Arg	Arg	Leu	Pro	Arg	Ser	Lys	Asp
	370				•	375					380			•	
Ala	Ile	Leu	Leu	Ala	Asp	Leu	Val	Asp	Ser	Cys	Asn	Ala	Gly	Asp	Glu
385	٠				390					395					400
T1e	Glu	Lan	Thr	Gl v	Tle	Tur	Hic	Asn	Acr	Tur	Asn	G1 v	Sor	Lan	Acr

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Thr	Ala	Asn	Gly	Phe	Pro	Val	Phe	Ala	Thr	Val	Ile	Leu	Ala	Asn	His
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Asp	Val	Lys	Met	Ile	Thr	Ser	Leu	Ser	Lys	Asp	Gln	Gln	Ile	Gly	Glu
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Lys	Ile	Phe	Ala	Ser	Ile	Ala	Pro	Ser	Ile	Tyr	Gly	His	Glu	Asp	Ile
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Lys	Arg	Gly	Pro	Ala	Leu	Ala	Leu	Phe	Gly	Gly	Glu	Pro	Lys	Asn	Pro
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Ser	Ser	Arg	Ala	Ile	Phe	Thr	Thr	Gly	Gln	Gly	Ala	Ser	Ala	Val	Ala
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Val	Thr	Ala	Tyr	Val	Gln	Arg	His	Pro	Val	Ser	Arg	Glu	Trp	Thr	Leu
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Glu	Ala	Gly	Ala	Leu	Val	Leu	Ala	Asp	Arg	Gly	Val	Cys	Leu	Ile	Asp
				565					570					575	
Glu	Phe	Asp	Lys	Met	Asn	Asp	G1n	Asp	Arg	Thr	Ser	Ile	His	Glu	Ala
	-		580	-				585					590		
Met	Glu	Gln	Gln	Ser	Ile	Ser	Ile	Ser	Lys	Ala	Ġly	Ile	Val	Thr	Ser
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Leu	Gln	Ala	Arg	Cys	Thr	Val	Ile	Ala	Ala	Ala	Asn	Pro	Ile	Ģly	Gly

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Arg	Tyr	Asp	Pro	Ser.	Leu	Thr	Phe	Ser	Glu	Asn	Val	Asp	Leu	Thr	Glu
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Asp	Pro	Val	Gln	Asp	Glu	Met	Leu	Ala	Arg	Phe	Val	Val	Gly	Ser	His
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Val	Arg	His	His	Pro	Ser	Asn	Lys	Glu	Glu	Glu	Gly	Leu	Ala	Asn	Gly
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Ser	Ala	Ala	Glu	Pro	Ala	Met	Pro	Asn	Thr	Tyr	Gly	Val	Glu	Pro	Leu
	690					695					700				
Pro	Gln.	Glu	Val	Leu	Lys	Lys	Tyr	Ile	Ile	Tyr	Ala	Lys	Glu	Arg	Val
705					710					715					720
His	Pro	Lys	Leu	Asn	Gln	Met	Asp	Gln	Asp	Lys	Val	Ala	Lys	Met	Tyr
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Ser	Asp	Leu	Arg	Lys	Glu	Ser	Met	Ala	Thr	Gly	Ser	Ile	Pro	Ile	Thr
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Val	Arg	His	Ile	Glu	Ser	Met	Ser	His	Gly	Gly	Gly	Pro	Arg	Ala	His
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Pro	Ser	Ala	Gly	Leu	Cys	Asp	Arg	Arg	Arg	Arg	Gln	His	Gly	His	Pro
	770				٠.	775					780				
Arg	Asp	Ala	Gly	Glu	Leu	His	Arg	His	Thr	Glu	Val	G1n	Arg	His	Arg
785	•				790		·			795					800
Ser	Met	Arg	Lys	Thr	Phe	Ala	Arg	Tyr	Leu	Ser	Phe	Arg	Arg	Asp	Asn
				805					810					815	
Asn	Glu	Leu	Leu	Leu	Phe	Ile	Leu	Lvs	Gln	Leu	Val	Ala	G1u	Gln	Val

830

96

820 825

Thr Tyr Gln Arg Asn Arg Phe Gly Ala Gln Gln Asp Thr Ile Glu Val
835 840 845

Pro Glu Lys Asp Leu Val Asp Lys Ala Arg Gln Ile Asn Ile His Asn 850 855 860

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i	aga	cgt	gct	gac	gcc	ctg	acc	tcc	agc	cct	ggc	aga	gac	ctc	ccc	cca	144
	Arg	Arg	Ala	Asp	Ala	Leu	Thr	Ser	Ser	Pro	Gly	Arg	Asp	Leu	Pro	Pro	
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			35					40					45				
	ttt	gaa	gat	gag	tct	gag	ggg	ctt	ctg	ggc	aca	gag	ggg	ссс	atg	gag	192
J	rne	Glu	ASP	GIU	ser	Glu	GIA	Leu	Leu	GIA	inr	GIU	GLY	Pro	мет	GIU	
		50					55					60					
-	gaa	gaa	gag	gat	gga	gag	gaa	ctc	att	ggt	gat	ggc	atg	gag	aga	gac	240
(	Glu	Glu	Glu	Asp	Gly	Glu	Glu	Leu	Ile	Gly	Asp	Gly	Met	Glu	Arg	Asp	
	65					70				•	75					80	
						. •					. 3					00	

tac cgt ccc att ccg gag ctc gat gtc tac gag gcc gag gga ttg gcc 288

Tyr Arg Pro Ile Pro Glu Leu Asp Val Tyr Glu Ala Glu Gly Leu Ala

85 90 95

ctg gat gat gaa gat gtg gag gag ctg aca gcc agt cag aga gag gca 336 Leu Asp Asp Glu Asp Val Glu Glu Leu Thr Ala Ser Gln Arg Glu Ala 100 105 110

gct gag cgg acc atg agg cag cgg gac cgt gag gct ggc aga ggc ctg 384

Ala Glu Arg Thr Met Arg Gln Arg Asp Arg Glu Ala Gly Arg Gly Leu

115 120 125

gga	cgc	atg	cgc	cgg.	ggg	ctg	ctc	tat	gac	agc	agc	gag	gaa	gat	gag	432
Gly	Arg	Met	Arg	Arg	Gly	Leu	Leu	Tyr	Asp	Ser	Ser	Glu	Glu	Asp	Glu	
	130					135					140					
gag	cgg	cct	gcc	cgt	aag	cgc	cgc	cac	gta	gaa	cgc	gcc	aca	gag	gat	480
Glu	Arg	Pro	Ala	Arg	Lys	Arg	Arg	His	Val	Glu	Arg	Ala	Thr	Glu	Asp	
145					150					155					160	
ggc	gag	gag	gat	gaa	gag	atg	atc	gag	agt	att	gag	aat	ctg	gag	gac	528
Gly	Glu	Glu	Asp	Glu	Glu	Met	Ile	Glu	Ser	Ile	Glu	Asn	Leu	Glu	Asp	
				165					170					175		
ctc	aag	ggc	cac	tcg	gtg	cgc	gag	cgg	gtg	agc	atg	gca	ggg	ссс	agg	576
Leu	Lys	Gly	His	Ser	Val	Arg	Glu	Arg	Val	Ser	Met	Ala	Gly	Pro	Arg	
			180					185					190			
ctg	gag	atc	cac	cac	cgc	ttc	aag	aac	ttc	ctg	cgc	acc	cac	gtg	gac	624
Leu	Glu	Ile	His	His	Arg	Phe	Lys	Asn	Phe	Leu	Arg	Thr	His	Val	Asp	
		195					200					205				
•								-								
agc	cat	ggc	cac	aac	gtc	ttc	aag	gag	cgc	atc	agt	gat	atg	tgc	aaa	672
Ser	His	Gly	His	Asn	Val	Phe	Lys	G1u	Arg	Ile	Ser	Asp	Met	Cys	Lys	
٠.	210					215					220					
							•									

Glu	Asn	Arg	Glu	Ser	Leu	Val	Val	Asn	Tyr	Glu	Asp	Leu	Ala	Ala	Arg	
225					230					235					240	
~n~	202	at a	++~	<b>700</b>	+00	++0	a t a		~~~	~~~		go+	~~~	++~	-+-	760
					tac											768
Glu	His	Val	Leu	Ala	Tyr	Phe	Leu	Pro	Glu	Ala	Pro	Ala	Glu	Leu	Leu	
				245					250					255		
cag	atc	ttt	gac	gag	gct	gcc	ctg	gag	gtc	gtg	ttg	gcc	atg	tac	cct	816
Gln	Ile	Phe	Asp	Glu	Ala	Ala	Leu	Glu	Val	Val	Leu	Ala	Met	Tyr	Pro	
			260					265					270			
222	tat	aac	cat	ato	200	220	cac	ato	cat	ata	c ac.	ato	too	020	ctg	864
															•	864
Lys	Tyr	Asp	Arg	He	Thr	Asn	His	lle	His	Val	Arg	lle	Ser	His	Leu	
		275				•	280					285				
cct	ctg	gtg	gag	gag	ctg	cgt	tca	ctg.	agg	cag	ttg	cac	ctg	aac	cag	912
Pro	Leu	Val	G1u	Glu	Leu	Arg	Ser	Leu	Arg	Gln	Leu	His	Leu	Asn	Gln	
	290					295					300					
					٠											
a+~		aat		o.a.t		~+~				+~~						060
					ggc						• 🗥		Λ.			960
Leu	Ile	Arg	Thr	Ser	Gly	Val	Val	Thr	Ser	Cys	Thr	Gly	Val	Leu	Pro	
305				:	310					315					320	
cag	ctc	agc	atg	gtc	aag	tac	aac	tgt	agc	aag	tgc	aac	ttt	gta	ctg	1008
Gĺn	Leu	Ser	Met	Val	Lys	Tyr	Asn	Cys	Ser	Lys	Cys	Asn	Phe	Val	Leu	
•					-	-		-		-	-					

ggg	cct	ttc	tgc	cag.	tct	cag	aat	cag	gag	gtg	aag	cct	ggc	tcc	tgc	1056
Gly	Pro	Phe	Cys	Gln	Ser	Gln	Asn	Gln	Glu	Val	Lys	Pro	Gly	Ser	Cys	
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cct	gag	tgc	cag	tct	gct	ggg	ccc	ttt	gag	atc	aac	atg	gag	gag	acc	1104
Pro	Glu	Cys	Gln	Ser	Ala	Gly	Pro	Phe	Glu	Ile	Asn	Met	Glu	Glu	Thr	
		355					360					365				
atc	tat	cag	aac	tac	caa	cgt	atc	cgc	atc	cag	gag	agt	ссс	ggc	aag	1152
Ile	Tyr	Gln	Asn	Tyr	Gln	Arg	Ile	Arg	Ile	Gln	Glu	Ser	Pro	G1y	Lys	
	370					375					380					
gtg	gcg	gct	ggc	cga	ctg	ccc	cgt	tcc	aag	gat	gcc	att	ctc	ctc	gct	1200
Val	Ala	Ala	Gly	Arg	Leu	Pro	Arg	Ser	Lys	Asp	Ala	Ile	Leu	Leu	Ala	
385					390					395					400	
gat	ctg	gtg	gac	agc	tgc	aag	cca	ggg	gac	gag	att	gag	ctg	acc	ggc	1248
Asp	Leu	Val	Asp	Ser	Cys	Lys	Pro	Gly	Asp	Glu	Ile	Glu	Leu	Thr	Gly	
				405					410					415		
											•					
att	tac	cat	aat	aac	tat	gac	ggc	tcg	ctt	aac	acc	gcc	aac	ggc	ttt	1296
					Tyr											
•	•		420		•			425					430			
-										÷ .						•

cca gtc ttt gcc act att atc ttg gcc aac cat gtt gcc aag aag gac 1344

Pro	Val	Phe	Ala	Thr	Ile	Ile	Leu	Ala	Asn	His	Val	Ala	Lys	Lys	Asp	
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aac	aaa	gta	gct	gtg	ggg	gag	ctc	acc	gat	gag	gac	gtg	aag	atg	atc	1392
Asn	Lys	Val	Ala	Val	Gly	Glu	Leu	Thr	Asp	Glu	Asp	Val	Lys	Met	Ile	
	450					455			٠		460					
acc	ggt	ctc	tcc	aag	gat	cag	caa	att	gga	gag	aag	atc	ttt	gcc	agc .	1440
Thr	Gly	Leu	Ser	Lys	Asp	Gln	Gln	Ile	Gly	Glu	Lys	Ile	Phe	Ala	Ser	
465					470					475					480	
att	gca	ccc	tcc	atc	·tat	ggg	cat	gaa	gac	atc	aag	aga	ggc	ctg	gct	1488
Ile	Ala	Pro	Ser	Ile	Tyr	Gly	His	Glu	Asp	Ile	Lys	Arg	Gly	Leu	Ala	
				485					490	٠.				495		
ctg	gcc	ctg	ttt	gga	ggg	gag	ccc	aag	aac	cca	ggt	gga	aag	cac	aag	1536
Leu	Ala	Leu	Phe	Gly	Gly	Glu	Pro	Lys	Asn	Pro	Gly	Gly	Lys	His	Lys	
			500					505					510			
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gtt	cga	ggt	gac	att	aat	gtg	ctc	ttg	tgt.	ggg	gac	cct	ggc	aca	gca	1584
Val	Arg	Gly	Asp	Ile	Asn	Val	Leu	Leu	Cys	Gly	Asp	Pro	Gly	Thr	Ala	
		515					520					525				
										9						
220	taa	000	tto	oto	222	taa	ata	an a		at a	tot	0.00	oat	<b>~~</b>	oto	1622

Lys Ser Gln Phe Leu Lys Tyr Ile Glu Lys Val Ser Ser Arg Ala Ile

540

535

ttc	acc	act	ggc	cag.	ggt	gcg	tca	gca	gtg	ggt	ctc	acc	gcg	tac	gtt	1680
Phe	Thr	Thr	Gly	G1n	Gly	Ala	Ser	Ala	Val	Gly	Leu	Thr	Ala	Tyr	Val	
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cag	cgg	cat	ссс	gtc	agc	aga	gag	tgg	acc	tta	gag	gcg	gga	gcc	ctg	1728
G1n	Arg	His	Pro	Val	Ser	Arg	Glu	Trp	Thr	Leu	Glu	Ala	Gly	Ala	Leu	
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gtt	ctg	gct	gac	cgg	ggg	gtg	tet	ctc	att	gac	gag	ttt	gac	aag	atg	1776
														Lys		
	Deu	nia	580	nu 5	·GIy	,41	O) S	585	110	пор	GIU	THE	590	ЦуЗ	inc t	
			360					363					550			
aat	gac	cag	gac	agg	acc	agc	atc	cac	gag	gcc	atg	gaa	cag	caa	agc	1824
Asn	Asp	Gln	Asp	Arg	Thr	Ser	Ile	His	Glu	Ala	Met	Glu	Gln	Gln	Ser	
		595					600					605				
							-									
atc	tcc	atc	tcc	aag	gct	ggc	atc	gtt	acc	tcg	ctg	caa	gcc	cgc	tgc	1872
Ile	Ser	Ile	Ser	Lys	Ala	Gly	Ile	Val	Thr	Ser	Leu	Gln	Ala	Arg	Cys	
	610					615		-			620					
act	gtc	ata	gct	gct	gcc	aac	ccc	ata	gga	ggc	cgc	tac	gac	cct	tca	1920
Thr	Val	Ile	Ala	Ala	Ala	Asn	Pro	Ile	Gly	Gly	Arg	Tyr	Asp	Pro	Ser	
625					630		•			635					640	
							•									
•																

ctg acc ttc tca gag aat gta gac ctc aca gag ccc atc att tcc cgc

Leu	Thr	Phe	Ser	Glu	Asn	Val	Asp	Leu	Thr	Glu	Pro	Ile	Ile	Ser	Arg
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:			660					665					670			
Phe	Asp	Val	Leu	Cys	Val	Val	Arg	Asp	Thr	Val	Asp	Pro	Val	Gln	Asp	
ttt	gat	gtc	ctg	tgt	gtg	gtg	agg	gac	act	gtt	gat	cca	gtt	cag	gat	2016

gag	atg	ctg	gcc	cgc	ttt	gtg	gtt	ggc	agc	cac	gtc	aga	cac	cac	ccc	2064
Glu	Met	Leu	Ala	Arg	Phe	Val	Val	Gly	Ser	His	Val	Arg	His	His	Pro	
		675				•	680					685				

agt	aac	aag	aag	gat	gaa	ggg	ttg	act	aat	ggt	ggc	acc	ttg	gag	cca	2112
Ser	Asn	Lys	Lys	Asp	Glu	Gly	Leu	Thr	Asn	Gly	Gly	Thr	Leu	Glu	Pro	•
	690					695					700					

gcç	atg	ccc	aac	aca	tat	ggc	gtg	gag	ccc	ctg	cct	cag	gag	gtg	ctg	2160
Ala	Met	Pro	Asn	Thr	Tyr	Gly	Val	Glu	Pro	Leu	Pro	Gln	Glu	Val	Leu	
705					710					715					720	-

aag	aag	tat	atc	atc	tat	gcc	aag	gag	agg	gtc	cgc	ccg	aag	ctc	aac	2208
Lys	Lys	Tyr	Ile	Ile	Tyr	Ala	Lys	Glu	Arg	Val	Arg	Pro	Lys	Leu	Asn	
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Glu	Ser	Met	Ala	Thr	Gly	Ser	Ile	Pro	Ile	Thr	Val	Arg	His	Ile	Glu	
		755					760					765				
tcc	atg	atc	cgc	atg	gcc	gag	gcc	cat	gcc	cgc	atg	cac	ctg	cgg	gac	2352
Ser	Met	Ile	Arg	Met	Ala	Glu	Ala	His	Ala	Arg	Met	His	Leu	Arg	Asp	
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Tyr	Val	Met	Glu	Asp	Asp	Val	Asn	Met	Ala	Ile	Arg	Val	Met	Met	Glu	
785					790					795					800	
agc	ttc	att	gac	acc	cag	aag	ttc	agc	gtc	atg	cgg	agt	atg	cgc	aag	2448
							Phe									
			-	805					810					815		
•			•							-				•		
act	ttt	gcc	Caa	tat	ctc	tcc	ttc	Cgg	cga	gat.	aac	aat	gat	ctg	ctg	2496
							Phe									2100
		7114	820	. , .	Leu	DCI		825	1118	пър	11511	non	830	Deu	Dou	
	*		020					020					000			
		_ 4													-	0544
							gtg								•	2544
Leu	Phe			Lys	GIn	Leu	Val	Ala	Glu	Gln	Val		Tyr	G1n	Arg	
		835					840					845				

aac cgc ttt ggg gcc cag cag gac acc att gaa ata cct gag aag gat

Asn Arg Phe Gly Ala Gln Gln Asp Thr Ile Glu Ile Pro Glu Lys Asp
850 855 860

ctg atg gac aag gcc agg cag atc aat att cac aac ctc tct gcc ttc 2640 Leu Met Asp Lys Ala Arg Gln Ile Asn Ile His Asn Leu Ser Ala Phe 865 870 875 880

tac gac agc gac ctc ttc aaa ttc aac aag ttc agc cgt gac ctg aaa 2688

Tyr Asp Ser Asp Leu Phe Lys Phe Asn Lys Phe Ser Arg Asp Leu Lys

885 890 895

cgc aaa ctg atc cta cag cag ttc tga 2715

Arg Lys Leu Ile Leu Gln Gln Phe
900 905

<210> 5

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<212> PRT

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**<400> 5** 

Met Ala Glu Ser Ser Glu Ser Leu Ser Ala Ser Ser Pro Ala Arg Gln

1 5 10 15

Arg Arg Ile Ser Asp Pro Leu Thr Ser Ser Pro Gly Arg Ser Ser

25

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30

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Glu	Glu	Glu	Asp	Gly	Glu	Glu	Leu	Ile	Gly	Asp	Gly	Met	Glu	Arg	Asp
65					70					75					80
Tyr	Arg	Pro	Ile	Pro	Glu	Leu	Asp	Val	Tyr	Glu	Ala	Glu	Gly	Leu	Ala
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Gly	Arg	Met	Arg	Arg	Gly	Leu	Leu	Tyr	Asp	Ser	Ser	Glu	Glu	Asp	Glu
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145					150					155					160
Gly	Gl <sub>u</sub>	Glu	Asp	Glu	Glu	Met	Ile	Glu	Ser	Ile	Glu	Asn	Leu	Glu	Asp
٠				165					170				٠.	175	
Leu	Lys	Gly	His	Ser	Val	Arg	Glu	Arg	Val	Ser	Met	Ala	Gly	Pro	Arg
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Leu	Glu	Ile	His	His	Arg	Phe	Lys	Asn	Phe	Leu	Arg	Thr	His	Val	Asp
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Ser	His	Gly	His	Asn	Val	Phe	Lys	Glu	Arg	Ile	Ser	Asp	Met	Cys	Lys
•	210					215	,				220				٠.
Gĺu	Asn	Arg	Glu	Ser	Leu	Val	Val	Asn	Tyr	Glu	Asp	Leu	Ala	Ala	Arg
225					220					225					240

Glu	His	Val	Leu	Ala	Tyr	Phe	Leu	Pro	Glu	Ala	Pro	Ala	Glu	Leu	Leu
				245.					250					255	
Gln	Ile	Phe	Asp	Glu	Ala	Ala	Leu	Glu	Val	Val	Leu	Ala	Met	Tyr	Pro
			260	•				265					270		
Lys	Tyr	Asp	Arg	Ile	Thr	Asn	His	Ile	His	Val	Arg	Ile	Ser	His	Leu
		275				**	280					285			·
Pro	Leu	Val	Glu	Glu	Leu	Arg	Ser	Leu	Arg	G1n	Leu	His	Leu	Asn	G1n
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Leu	Ile	Arg	Thr	Ser	Gly	Val	Val	Thr	Ser	Cys	Thr	Gly	Val	Leu	Pro
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Gln	Leu	Ser	Met	Val	Lys	Tyr	Asn	Cys	Ser	Lys	Cys	Asn	Phe	Val	Leu
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Ile	Tyr	Gln	Asn	Tyr	G1n	Arg	Ile	Arg	Ile	Gln	Glu	Ser	Pro	Gly	Lys
	370	)				375					380		٠.		
Val	. Ala	Ala	Gly	Arg	Leu	Pro	Arg	Ser	Lys	Asp	Ala	Ile	Leu	Leu	Ala
385	5				390	١	٠			395					400
Asp	Leu	ı Val	Asp	Ser	Cys	Lys	Pro	Gly	Asp	Glu	Ile	Glu	Leu	Thr	Gly
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Ιlϵ	e Tyr	His	s Asn	Asn	Tyr	Asp	Gly	Ser	Leu	Asn	Thr	Ala	Asn	Gly	Phe
•	.*		420	)				425	;				430		
Pro	Val	Phe	e Ala	Thr	· Ile	Ile	e Leu	Ala	Asn	His	Val	Ala	Lys	Lys	Asp
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Asn	Lys	Val	Ala	Val	Gly	Glu	Leu	Thr	Asp	Glu	Asp	Val	Lys	Met	Ile
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Thr	Gly	Leu	Ser	Lys	Asp	Gln	Gln	Ile	Gly	Glu	Lys	Ile	Phe	Ala	Ser
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Ile	Ala	Pro	Ser	Ile	Tyr	Gly	His	Glu	Asp	Ile	Lys	Arg	Gly	Leu	Ala
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Val	Arg	Gly	Asp	Ile	Asn	Val	Leu	Leu	Cys	Gly	Asp	Pro	Gly	Thr	Ala
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Gln	Arg	His	Pro	Val	Ser	Arg	Glu	Trp	Thr	Leu	Glu	Ala	Gly	Ala	Leu
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Val	Leu	Ala	Asp	Arg	Gly	Val	Cys	Leu	Ile	Asp	Glu	Phe	Asp	Lys	Met
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Ile	Ser	Ile	Ser	Lys	Ala	Gly	Ile	Val	Thr	Ser	Leu	Gln	Ala	Arg	Cys
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Leu	Thr	Phe	Ser	Glu	Asn	Val	Asp	Leu	Thr	Glu	Pro	Ile	Ile	Ser	Arg
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Phe	Asp	Val	Leu	Cys	Val	Val	Arg	Asp	Thr	Val	Asp	Pro	Val	Gln	Asp
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Glu	Met	Leu	Ala	Arg	Phe	Val	Val	Gly	Ser	His	Val	Arg	His	His	Pro
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Ser	Asn	Lys	Lys	Asp	Glu	Gly	Leu	Thr	Asn	Gly	Gly	Thr	Leu	Glu	Pro
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Ala	Met	Pro	Asn	Thr	Tyr	Gly	Val	Glu	Pro	Leu	Pro	Gln	Glu	Val	Leu
705		ė			710					715					720
Lys	Lys	Tyr	Ile	Ile	Tyr	Ala	Lys	Glu	Arg	Val	Arg	Pro	Lys	Leu	Asn
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Gln	Met	Asp	Gln	Asp	Lys	Val	Ala	Arg	Met	Tyr	Ser	Asp	Leu	Arg	Lys
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Glu	Ser	Met	Ala	Thr	Gly	Ser	Ile	Pro	Ile	Thr	Val	Arg	His	Ile	Glu
•		755					760					765			
Ser	Met	Ile	Arg	Met	Ala	Glu	Ala	His	Ala	Arg	Met	His	Leu	Arg	Asp
	770					775					780				
Tyr	Val	Met	Glu	Asp	Asp	Val	Asn	Met	Ala	Ile	Arg	Val	Met	Met	Glu
785				*	790					795	*				800
Ser	Phe	Ile	Asp	Thr	G1n	Lys	Phe	Ser	Val	Met	Arg	Ser	Met	Arg	Lys
			٠	805					810					815	
Thr	Phe	Ala	Arg	Tyr	Leu	Ser	Phe	Arg	Arg	Asp	Asn	Asn	Asp	Leu	Leu
•	·. ·	.•	- 820	)				825					830	•	
Leu	Phe	.Ile	Leu	Lys	Gln	Leu	Val	Ala	· Glu	Gln	Val	Thr	Tyr	Gln	Arg
:		835	<b>5</b>				840	1				845	5		
Asn	Arg	Phe	e Gly	Ala	Gln	Gln	Asp	Thr	·Ile	Glu	ı Ile	Pro	Glu	l Lys	Asp
	ÖEA					822					860	)			

Leu Met Asp Lys Ala Arg Gln Ile Asn Ile His Asn Leu Ser Ala Phe 865 870 875 880

Tyr Asp Ser Asp Leu Phe Lys Phe Asn Lys Phe Ser Arg Asp Leu Lys

885

890

895

Arg Lys Leu Ile Leu Gln Gln Phe
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<212> DNA

<213> Homo sapiens

<220>

<221> CDS

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aattggcttg tg atg gag gcg tct ttg ggg att cag atg gat gag cca atg 171 Met Glu Ala Ser Leu Gly Ile Gln Met Asp Glu Pro Met

1

5

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а	aa	aac	gag	cag	aat	ttt	aaa	ctt	gca	ggt	gtt	aaa	aaa	gat	att	gag	267
L	ys	Asn	Glu	Gln	Asn	Phe	Lys	Leu	Ala	Gly	Val	Lys	Lys	Åsp	Ile	Glu	
	30					35	-				40					45	
		-										•					
a	iag	ctt	tat	gaa	gct	gta	cca	cag	ctt	agt	aat	gtg	ttt	aag	att	gag	315
							Pro							•			
					50					55					60		
•	ac	aaa	att	gga	gaa	ggc	act	ttc	agc	tct	gtt	tat	ttg	gcc	aca	gca	363
							Thr										
1	rsp	Буз	110	65		OI,	••••		70			,					
											•			•			
	•									. 4. 4		- 4			++~	o++	411
							gaa										411
(	Gln	Leu	G1n	Val	Gly	Pro	Glu	Glu	Lys	lle	Ala	Leu	Lys	His	Leu	. 11e	
		. •	80					85					90				
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	cca	aca	agt	cat	cct	ata	aga	att	gca	gct	gaa	ctt	cag	tgc	cta	aca	459
·. ]	Pro	Thr	Ser	His	Pro	Ile	Arg	Ile	Ala	Ala	Glu	Leu	Gln	Cys	Leu	Thr	
		95	;			·	100		•			105	5				
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	Val	Ala	Gly	Gly	Gln	. Asp	Asn	Val	Met	Gly	Val	Lys	з Туг	Cys	: Phe	. Arg	

aag	aat	gat	cat	gta.	gtt	att	gct	atg	cca	tat	ctg	gag	cat	gag	tcg	555
Lys	Asn	Asp	His	Val	Val	Ile	Ala	Met	Pro	Tyr	Leu	Glu	His	Glu	Ser	•
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		-														•
ttt	ttg	gac	att	ctg	aat	tct	ctt	tcc	ttt	caa	gaa	gta	cgg	gaa	tat	603
Phe	Leu	Asp	Ile	Leu	Asn	Ser	Leu	Ser	Phe	Gln	Glu	Val	Arg	Glu	Tyr	
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atg	ctt	aat	ctg	ttc	aaa	gct	ttg	aaa	cgc	att	cat	cag	ttt	ggt	att	651
												Gln				
1.100	200	160					165					170				
		100														÷
		o or t	got	at t	220	ccc	age	aat	tit	t.t.a	tat	aat	agg	cgc	ctg	699
												Asn				
val			Asp	vai	Lys			ASII	THE	LCu	185		1111 8		20-	•
•	175					180	ı				100	ı				
																747
															gat	747
Lys	Lys	Tyr	Ala	Leu	Val	Asp	Phe	Gly	Leu	Ala	Glr	Gly	Thr	His	Asp	
190					195	5				200	)				205	
					. :										*	•
acg	aaa	ata	a gag	g ctt	ctt	aaa	ttt	gto	cag	g tct	gaa	a gct	cag	g cag	gaa	795
Thr	Lys	s Ile	e Glu	ı Let	ı Let	ı Lys	Phe	e Val	Glr	ı Sei	c Glu	ı Ala	Glr	ı Glr	Glu	• .
				210	)				215	5				220		•

agg tgt tca caa aac aaa tcc cac ata atc aca gga aac aag att cca

								J 4.	/ 0	3						
Arg	Cys	Ser	Gln	Asn	Lys	Ser	His	Ile	Ile	Thr	Gly	Asn	Lys	Ile	Pro	:
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Leu	Ser	Gly	Pro	Val	Pro	Lys	Glu	Leu	Asp	Gln	Gln	Ser	Thr	Thr	Lys	
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Ala	Ser	Val	Lys	Arg	Pro	Tyr	Thr	Asn	Ala	Gln	Ile	Gln	Ile	Lys	G1n	
	255					260					265					
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Gly	Lys	Asp	Gly	Lys	Glu	Gly	Ser	Val	Gly	Leu	Ser	Va]	l Glm	Arg	g Ser	
270	)				275					280	) · .				285	
gtt	t tt1	gga	gaa	a aga	aat	tto	aat	ata	cac	ago	tcc	at	t tca	a cat	t gag	1035
Va]	l Phe	e Gly	g Glu	ı Arg	g Asn	Phe	e Asn	Ile	His	s Ser	Ser	· Ile	e Sei	r His	s Glu	
				290	)				295	5				306	0	
								٠.								
ago	c cc	t gc	a gt	g aaa	a cto	ate	g aag	cag	g tca	a aag	g act	t gt	g ga	t gt	a ctg	1083
Sea	r Pro	o Ala	a Va	l Lys	s Lei	ı Met	t Lys	G1r	s Sei	r Lys	s Thi	r Va	1 As	p Va	l Leu	
			30	5	٠.			310	)				31	5		
												٠.		•		
to	t. ао	์ ล ลล	g tt:	a gc	a aca	a aaa	a aag	g aag	g gc	t at	t tc	t ac	g aa	a gt	t_atg	1131

Ser Arg Lys Leu Ala Thr Lys Lys Lys Ala Ile Ser Thr Lys Val Met

aat	agt	gct	gtg	atg.	agg	aaa	act	gcc	agt	tct	tgc	cca	gct	agc	ctg	1179
Asn	Ser	Ala	Val	Met	Arg	Lys	Thr	Ala	Ser	Ser	Cys	Pro	Ala	Ser	Leu	
	335					340	٠				345				٠.	
									•							
acc	tgt	gac	tgc	tat	gca	aca	gat	aaa	gtt	tgt	agt	att	tgc	ctt	tca	1227
Thr	Cys	Asp	Cys	Tyr	Ala	Thr	Asp	Lys	Val	Cys	Ser	Ile	Cys	Leu	Ser	
350					355					360		••			365	
	•														·	
agg	cgt	cag	cag	gtt	gcc	cct	agg	gca	ggt	aca	cca	gga	ttc	aga	gca	1275
Arg	Arg	Gln	G1n	Val	Ala	Pro	Arg	Ala	Gly	Thr	Pro	Gly	Phe	Arg	Ala	
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cca	gag	gtc	ttg	aca	aag	tgc	ссс	aat	caa	act	aca	gca	att	gac	atg	1323
Pro	Glu	Val	Leu	Thr	Lys	Cys	Pro	Asn	Gln	Thr	Thr	Ala	Ile	Asp	Met	
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:	415					420		,			425		•			
	110	•														

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Asp	His	Lys	Ala	Ser	Cys	Leu	Val	Gln	Thr	Pro	Pro	Gly	Gln	Tyr	Ser	
	495	;	_		•	500					505					
						; · ·										
ggg	aat	tca	ı ttt	aaa	aag	ggg	gat	agt	aat	agc	tgt	gag	cat	tgt	ttt	1707
•																
Glv							Asp	Ser		Ser		Glu	His		s Phe	,
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gct tat gac ctg ctt gat aaa ctt cta gat cta aat cca gct tca aga 1803

Ala Tyr Asp Leu Leu Asp Lys Leu Leu Asp Leu Asp Pro Ala Ser Arg

545 550 555

ata aca gca gaa gaa gct ttg ttg cat cca ttt ttt aaa gat atg agc 1851

Ile Thr Ala Glu Glu Ala Leu Leu His Pro Phe Phe Lys Asp Met Ser

560 565 570

ttg tga taatggatct tcatttaatg tttactgtta tgaggtagaa taaaaaagaa 1907 Leu

575

tactttgtaa tagccacaag ttcttgttta gagaccagag caggattaat aatttatttt 1967
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<210> 7

<211> 574

<212> PRT

<213> Homo sapiens

<400> 7

Met Glu Ala Ser Leu Gly Ile Gln Met Asp Glu Pro Met Ala Phe Ser

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Pro Gln Arg Asp Arg Phe Gln Ala Glu Gly Ser Leu Lys Lys Asn Glu

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Gln Asn Phe Lys Leu Ala Gly Val Lys Lys Asp Ile Glu Lys Leu Tyr

35 40 45

Glu Ala Val Pro Gln Leu Ser Asn Val Phe Lys Ile Glu Asp Lys Ile

50 55 60

Gly Glu Gly Thr Phe Ser Ser Val Tyr Leu Ala Thr Ala Gln Leu Gln

65					70					75					80
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His	Pro	Ile	Arg	Ile	Ala	Ala	Glu	Leu	Gln	Cys	Leu	Thr	Val	Ala	Gly
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His	Val	Val	Ile	Ala	Met	Pro	Tyr	Leu	Glu	His	Glu	Ser	Phe	Leu	Asp
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Ala	Leu	Val	Asp	Phe	Gly	Leu	Ala	Gln	Gly	Thr	His	Asp	Thr	Lys	Ile
		195					200					205			
Glu	Leu	Leu	Lys	Phe	Val	Gln	Ser	Glu	Ala	G1n	Gln	Glu	Arg	Cys	Ser
	210					215					220				
Gln	Asn	Lys	Ser	His	Ile	Ile	Thr	Gly	Asn	Lys	Ile	Pro	Leu	Ser	Gly
-225			٠		230					235	;				240
Pro	Val	Pro	Lys	Glu	Leu	Asp	G1n	Gln	Ser	Thr	Thr	Lys	Ala	Ser	Val
	·			245	;				250					255	
Lys	Arg	Pro	Tyr	Thr	Asn	Ala	Gln	Ile	Gln	Ile	Lys	Gln	Gly	Lys	Asp
• •			260	)				265	•	•			270		
C1	· T	C1.	C1.		. Vol	61.	Lau	Sor	· Val	G1r	Aro	Ser	· Val	Phe	Glv

		275					280					285				
Glu	Arg	Asn	Phe	Asn.	Ile	His	Ser	Ser	Ile	Ser	His	Glu	Ser	Pro	Ala	
	290					295					300	•		<i>I</i>		
/al	Lys	Leu	Met	Lys	Gln	Ser	Lys	Thr	Val	Asp	Val	Leu	Ser	Arg	Lys	
305					310					315					320	
Leu	Ala	Thr	Lys	Lys	Lys	Ala	Įle	Ser	Thr	Lys	Val	Met	Asn	Ser	Ala	
				325		-			330					335		
Val	Met	Arg	Lys	Thr	Ala	Ser	Ser	Cys	Pro	Ala	Ser	Leu	Thr	Cys	Asp	
			340			•		345					350			
Cys	Tyr	Ala	Thr	Asp	Lys	Val	Cys	Ser	Ile	Cys	Leu	Ser	Arg	Arg	Gln	
		355					360					365				
Gln	Val	Ala	Pro	Arg	Ala	G1y	Thr	Pro	G1y	Phe	Arg	Ala	Pro	Glu	Val	
	370					375		٠			380					
Leu	Thr	Lys	Cys	Pro	Asn	Gln	Thr	Thr	Ala	Ile	Asp	Met	Trp	Ser	Ala	
385					390			-		395					400	
Gly	Val	Ile	Phe	Leu	Ser	Leu	Leu	Ser	Gly	Arg	Tyr	Pro	Phe	Tyr	Lys	
				405					410					415		
Ala	Ser	Asp	Asp	Leu	Thr	Ala	Leu	Ala	Gln	Ile	Met	Thr	Ile	Arg	Gly	
		•	420					425	•				430	)		
Ser	Arg	Glu	Thr	Ile	Gln	Ala	Ala	Lys	Thr	Phe	Gly	Lys	Ser	Ile	Leu	
	•	435	i				440					445	5			
Cys	Ser	Lys	Glu	Val	Pro	Ala	G1n	Asp	Leu	Arg	Lys	Leu	ı Cys	Glu	ı Arg	
	450	)			-	455					460	)				
Leu	Arg	Gly	Met	. Asp	Ser	Ser	Thr	Pro	Lys	Leu	1 Thr	Ser	Asp	Ile	Gln	ı
465					470					475	5				480	•
C1	и: -	. 41-	. 9	. His	. G1 =	Pro	Δ1-	Tle	Ser	- G1:	1 J.vs	The	- Asr	His	Lvs	

485 490 495

Ala Ser Cys Leu Val Gln Thr Pro Pro Gly Gln Tyr Ser Gly Asn Ser 500 505 510

Phe Lys Lys Gly Asp Ser Asn Ser Cys Glu His Cys Phe Asp Glu Tyr
515 520 525

Asn Thr Asn Leu Glu Gly Trp Asn Glu Val Pro Asp Glu Ala Tyr Asp
530 535 540

Leu Leu Asp Lys Leu Leu Asp Leu Asn Pro Ala Ser Arg Ile Thr Ala 545 550 555 560

Glu Glu Ala Leu Leu His Pro Phe Phe Lys Asp Met Ser Leu
565 570

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<212> DNA

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<220>

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<222> (518)..(2542)

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ggcaggcacg	aggggcgagc	gcgaggcggg	gcacggcgcg	tggcgtgaga	cggggcgggg	240
cgcgcgtatc	ggcgccgcgg	ccgcgtgacg	cgttttcaaa	tcttcaaccg	ccgcagccca	300
ctcgtttgtg	ctttgcgcct	tcctcctccg	cgccttggag	ccggatccgg	ccccggaaac	360
ccgacctgca	gacgcggtac	ctctactgcg	tagaggccgt	agctggcgga	aggagagagg	420
cggccgtcct	gtcaacaggc	cgggggaagc	cgtgctttcg	cggctgcccg	gtgcgacact	480

ttctccggac ccagcatgta ggtgccgggc gactgcc atg aac tcc gga gcc atg 535

Met Asn Ser Gly Ala Met

1 5

agg atc cac agt aaa gga cat ttc cag ggt gga atc caa gtc aaa aat 583
Arg Ile His Ser Lys Gly His Phe Gln Gly Gly Ile Gln Val Lys Asn
10 15 20

gaa aaa aac aga cca tct ctg aaa tct ctg aaa act gat aac agg cca 631
Glu Lys Asn Arg Pro Ser Leu Lys Ser Leu Lys Thr Asp Asn Arg Pro
25 30 35

gaa aaa tcc aaa tgt aag cca ctt tgg gga aaa gta ttt tac ctt gac 679

Glu Lys Ser Lys Cys Lys Pro Leu Trp Gly Lys Val Phe Tyr Leu Asp

40 45 50

tta cct tct gtc acc ata tct gaa aaa ctt caa aag gac att aag gat

727

Leu Pro Ser Val Thr Ile Ser Glu Lys Leu Gln Lys Asp Ile Lys Asp

55

60

65

70

ctg	gga	ggg	cga	gtt	gaa	gaa	ttt	ctc	agc	aaa	gat	atc	agt	tat	ctt	775
Leu	Gly	Gly	Arg	Val.	Glu	Glu	Phe	Leu	Ser	Lys	Asp	Ile	Ser	Tyr	Leu	
				75					80					. 85		
•																
att	tca	aat	aag	aag	gaa	gct	aaa	ttt	gca	caa	acc	ttg	ggt	cga	att	823
Ile	Ser	Asn	Lys	Lys	Glu	Ala	Lys	Phe	Ala	Gln	Thr	Leu	Gly	Arg	Ile	
			90					95					100			
													٠			
tct	cct	gta	cca	agt	cca	gaa	tct	gca	tat	act	gca	gaa	acc	act	tca	871
Ser	Pro	Val	Pro	Ser	Pro	Glu	Ser	Ala	Tyr	Thr	Ala	Glu	Thr	Thr	Ser	
		105					110					115				
cct	cat	ccc	agc	cat	gat	gga	agt	tca	ttt	aag	tca	cca	gac	aca	gtg	919
Pro	His	Pro	Ser	His	Asp	Gly	Ser	Ser	Phe	Lys	Ser	Pro	Asp	Thr	Val	
	120	)				125					130					
						-										
tgt	tta	ago	aga	gga	aaa	tta	tta	gtt	gaa	aaa	gct	ato	aag	gac	cat	967
Cys	Leu	ı Ser	Arg	Gly	Lys	Leu	Leu	Val	Glu	Lys	Ala	ı Ile	Lys	s Asp	His	
135	;			-	140	) .				145					150	
													-			
gat	tţt	ätt	cct	tca	a aat	agt	ata	tta	tca	aat	gco	tte	g tca	ı tgg	g gga	1015
											,	•			o Gly	
·				159					160					16		
. •														. •		

gta aaa att ctt cat att gat gac att aga tac tac att gaa caa aag

Val Lys Ile Leu His Ile Asp Asp Ile Arg Tyr Tyr Ile Glu Gln Lys

aaa	aaa	gag	ttg	tat	tta	ctc	aag	aaa	tca	agt	act	tca	gta	aga	gat	1111
Lys	Lys	Glu	Leu	Tyr	Leu	Leu	Lys	Lys	Ser	Ser	Thr	Ser	Val	Arg	Asp	
		185					190					195				
				•	-											
ggg	ggc	aaa	aga	gtt	ggt	agt	ggt	gca	caa	aaa	aca	aga	aca	gga	aga	1159
Gly	Gly	Lys	Arg	Val	Gly	Ser	Gly	Ala	Gln	Lys	Thr	Arg	Thr	Gly	Arg	
	200					205					210					
ctc	aaa	aag	cct	ttt	gta	aag	gtg	gaa	gat	atg	agc	caa	ctt	tat	agg	1207
Leu	Lys	Lys	Pro	Phe	Val	Lys	Val	Glu	Asp	Met	Ser	Gln	Leu	Tyr	Arg	
215					220					225					230	
cca	ttt	tat	ctt	cag	ctg	acc	aat	atg	cct	ttt	ata	aat	tat	tct	att	1255
															Ile	
				235					240					245		
cag	ลลฮ	ccc	tgc	agt.	cca	ttt	gat	gta	gac	aag	cca	tct	agt	atg	caa	1303
			٠				-								Gln	
OIII	Lys	110	250			1110		255		,			260			
•			200	,				200								
		·		· ·							. ast	gar	v gat	- 220	r tat	1351
aag	caa	act	cag	gtt	8 888	CLA	aga	all	· caa	aca	gal	650	, 801		tat	1001

Lys Gln Thr Gln Val Lys Leu Arg Ile Gln Thr Asp Gly Asp Lys Tyr

					,			40,	/ 0	5						
ggt	gga	acc	tca	att	caa	ctc	cag	ttg	aaa	gag	aag	aag	aaa	aaa	gga	1399
Gly	Gly	Thr	Ser	Ile.	Gln	Leu	Gln	Leu	Lys	Glu	Lys	Lys	Lys	Lys	Gly	
	280					285					290					
tat	tgt	gaa	tgt	tgc	ttg	cag	aaa	tat	gaa	gat	cta	gaa	act	cac	ctt	1447
							Lys									
295					300					305					310	
											٠					
cta	agt	gag	caa	cac	aga	aac	ttt	gca	cag	agt	aac	cag	tat	caa	gtt	1495
							Phe									
				315	-				320					325		
									•							
gtt	gat	gat	att	gta	tct	aag	tta	gtt	ttt	gac	ttt	gtg	gaa	tat	gaa	1543
															Glu	
		•	330		-			335					340			
									•							
ลลต	· ˈgac	aca	cct	: aaa	aag	aaa	aga	ata	aaa	tac	agt	gtt	gga	tco	ctt	1591
									-						Leu	
D) S	, HSD	345		. 2,5	, .	_,_	350					355			٠.	
		<b>J</b> 10	•	•				·								
+ 0 <del>1</del>	+	· ~++	. + 0 +	t gra	i aat	gto	cta	. ลลล	เลลต	act	gaa	caa	ı aas	g gaa	a aaa	1639
															ı Lys	
Ser			Se1	. WIS	ı ser			гр'nя	о Буз	, i <sub>111</sub>	370		,.	, 510		
	360	)				.365	)				316	,				

gtg gaa ttg caa cat att tct cag aaa gat tgc cag gaa gat gat aca 1687 Val Glu Leu Gln His Ile Ser Gln Lys Asp Cys Gln Glu Asp Asp Thr aca gtg aag gag cag aat ttc ctg tat aaa gag acc cag gaa act gaa 1735.

Thr Val Lys Glu Gln Asn Phe Leu Tyr Lys Glu Thr Gln Glu Thr Glu

395 400 405

aaa aag ctc ctg ttt att tca gag ccc atc ccc cac cct tca aat gaa 1783

Lys Lys Leu Leu Phe Ile Ser Glu Pro Ile Pro His Pro Ser Asn Glu

410 415 420

ttg aga ggg ctt aat gag aaa atg agt aat aaa tgt tcc atg tta agt 1831 Leu Arg Gly Leu Asn Glu Lys Met Ser Asn Lys Cys Ser Met Leu Ser 425 430 435

aca gct gaa gat gac ata aga cag aat ttt aca cag cta cct cta cat

1879

Thr Ala Glu Asp Asp Ile Arg Gln Asn Phe Thr Gln Leu Pro Leu His

440

445

450

aaa aac aaa cag gaa tgc att ctt gac att tcc gaa cac aca tta agt 1927
Lys Asn Lys Gln Glu Cys Ile Leu Asp Ile Ser Glu His Thr Leu Ser
455 460 465 470

gaa aat gac tta gaa gaa cta agg gta gat cac tat aaa tgt aac ata 1975 Glu Asn Asp Leu Glu Glu Leu Arg Val Asp His Tyr Lys Cys Asn Ile 475 480 485

									4 1	, 0	9						
С	ag	gca	tct	gta	cat	gtt	tct	gat	ttc	.agt	aca	gat	aat	agt	gga	tct	2023
G	ln	Ala	Ser	Val	His.	Val	Ser	Asp	Phe	Ser	Thr	Asp	Asn	Ser	Gly	Ser	•
				490					495					500			
c	aa	cca	aaa	cag	aag	tca	gat	act	gtg	ctt	tţt	cca	gca	aag	gat	ctc	2071
C	ln	Pro	Lys	Gln	Lys	Ser	Asp	Thr	Val	Leu	Phe	Pro	Ala	Lys	Asp	Leu	
			505			•		510					515				
												• *					
8	aag	gaa	aag	gac	ctt	cat	tca	ata	ttt	act	cat	gat	tct	ggt	ctg	ata	2119
I	_ys	Glu	Lys	Asp	Leu	His	Ser	Ile	Phe	Thr	His	Asp	Ser	Gly	Leu	Ile	
		520					525					530					
														•			
á	aca	ata	aac	agt	tca	caa	gag	cac	cta	act	gtt	cag	gca	aag	gct	cca	2167
•	Γhr	Ile	Asn	Ser	Ser	Gln	Glu	His	Leu	Thr	Val	Gln	Ala	Lys	Ala	Pro	
;	535					540					545					550	
												٠					
	ttc	cat	act	cct	cct	gag	gaa	ccc	aat	gaa	tgt	gac	ttc	aag	aat	atg	2215
ì	Phe	His	Thr	Pro	Pro	Glu	Glu	Pro	Asn	Glu	Cys	Asp	Phe	Lys	Asn	Met	
					555					560				-	565		
										-				-			
	gat	agt	tta	cct	tct	ggt	aaa	ata	cat	cga	aaa	gtg	aaa	ata	ata	tta	2263
	Asp	Ser	Leu	Pro	Ser	Gly	Lys	Ile	His	Arg	Lys	Val	Lys	: Ile	Ile	Leu	•

gga cga aat aga aaa gaa aat ctg gaa cca aat gct gaa ttt gat aaa 2311 Gly Arg Asn Arg Lys Glu Asn Leu Glu Pro Asn Ala Glu Phe Asp Lys

575

570

aga act gaa ttt att aca caa gaa gaa aac aga att tgt agt tca ccg Arg Thr Glu Phe Ile Thr Gln Glu Glu Asn Arg Ile Cys Ser Ser Pro gta cag tct tta cta gac ttg ttt cag act agt gaa gag aaa tca gaa Val Gln Ser Leu Leu Asp Leu Phe Gln Thr Ser Glu Glu Lys Ser Glu ttt ttg ggt ttc aca agc tac aca gaa aag agt ggt ata tgc aat gtt Phe Leu Gly Phe Thr Ser Tyr Thr Glu Lys Ser Gly Ile Cys Asn Val tta gat att tgg gaa gag gaa aat tca gat aat ctg tta aca gcg ttt Leu Asp Ile Trp Glu Glu Glu Asn Ser Asp Asn Leu Leu Thr Ala Phe ttc tcg tcc cct tca act tct aca ttt act ggc ttt tag aatttaaaaa Phe Ser Ser Pro Ser Thr Ser Thr Phe Thr Gly Phe

atgcatactt ttcagaagtg ataaggatca tattcttgaa atttttataa atatgtatgg 2612 aaattcttag gatttttta ccagctttgt ttacagaccc aaatgtaaat attaaaaata 2672 aatatttgca attttctaca gaattgaata cctgttaaag aaaaattaca gaataaactt 2732 gtgactggtc ttgttttaca ttaaaaaaaa aaaaaaaaa aactcgag 2780

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<211> 674 <212> PRT <213> Homo sapiens <400> 9 Met Asn Ser Gly Ala Met Arg Ile His Ser Lys Gly His Phe Gln Gly Gly Ile Gln Val Lys Asn Glu Lys Asn Arg Pro Ser Leu Lys Ser Leu Lys Thr Asp Asn Arg Pro Glu Lys Ser Lys Cys Lys Pro Leu Trp Gly Lys Val Phe Tyr Leu Asp Leu Pro Ser Val Thr Ile Ser Glu Lys Leu Gln Lys Asp Ile Lys Asp Leu Gly Gly Arg Val Glu Glu Phe Leu Ser Lys Asp Ile Ser Tyr Leu Ile Ser Asn Lys Lys Glu Ala Lys Phe Ala Gln Thr Leu Gly Arg Ile Ser Pro Val Pro Ser Pro Glu Ser Ala Tyr Thr Ala Glu Thr Thr Ser Pro His Pro Ser His Asp Gly Ser Ser Phe Lys Ser Pro Asp Thr Val Cys Leu Ser Arg Gly Lys Leu Leu Val Glu 

Ive	Ala	Πe	Lvs	Asp	His	Asp	Phe	Ile	Pro	Ser	Asn	Ser	Ile	Leu	Ser
	1114	110	2,3		150	P				155					160
145		_				17 3		T1.	T		T1 a	Aan	Aán	110	
Asn	Ala	Leu	Ser		Gly	Val	Lys	He		nis	11e	ASP	ASP		ni g
		•		165					170					175	•
Tyr	Tyr	Ile	Glu	Gln	Lys	Lys	Lys	Glu	Leu	Tyr	Leu	Leu	Lys	Lys	Ser
			180					185					190		
Ser	Thr	Ser	Val	Arg	Asp	Gly	Gly	Lys	Arg	Val	Gly	Ser	Gly	Ala	Gln
	•	195				•	200	-				205			
Lys	Thr	Arg	Thr	Gly	Arg	Leu	Lys	Lys	Pro	Phe	Val	Lys	Val	Glu	Asp
	210					215					220				
Met	Ser	Gln	Leu	Tyr	Arg	Pro	Phe	Tyr	Leu	G1n	Leu	Thr	Asn	Met	Pro
225					230					235					240
	Ile	Asn	Tyr	Ser	Ile	Gln	Lys	Pro	Cys	Ser	Pro	Phe	Asp	Val	Asp
			•	245					250					255	
Lve	Pro	Ser	Ser			Lvs	: Gln	Thr	Gln	Val	Lys	Leu	Arg	Ile	Gln
Lys	110	961			0111	. 2,	. 0111	265			J		270		
			260		· m	01	C1			. 11.	Cl <sub>n</sub>	Lou			Ive
Thr	Asp			Lys	lyr	GIY			. Set	. 116	GII			Leu	Lys
	-	275			•		280					285		_	2.3
G1u	Lys	Lys	Lys	Lys	Gly	Туз	Cys	Glu	ı Cys	Cys	Leu	ı Gln	ı Lys	; Tyr	Glu
	290	)				299	5				300	)			
Asp	Leu	ı Glu	ı Thr	His	s Leu	ı Let	ı Ser	Glu	ı Glr	n His	Are	g Asr	ı Ph∈	Ala	Gln
305	5				310	)				315	5 .				320
Ser	Asr	ı Glı	n Tyi	Glr	n Val	l Va	l Asp	Ası	o Ile	e Val	Sei	c Lys	s Lei	ı Val	Phe
•			-	329	5				330	) ·		•		335	5
Asr	Phe	e Vai	1 G11	1 Туз	r Glı	ı Ly:	s Ası	Th:	r Pro	o Lys	s Lys	s Lys	s Arg	g Ile	e Lys
•			211	1				34!					350		

Γyr	Ser	Val	Gly	Ser	Leu	Ser	Pro	Val	Ser	Ala	Ser	Val	Leu	Lys	Lys
		355					360					365			
Thr	Glu	G1n	Lys	Glu	Lys	Val	Glu	Leu	Gln	His	Ile	Ser	Gln	Lys	Asp
-	370					375		•			380				
Cys	Gln	G1u	Asp	Asp	Thr	Thr	Val	Lys	Glu	Gln	Asn	Phe	Leu	Tyr	Lys
385					390					395					400
Glu	Thr	Gln	Glu	Thr	Glu	Lys	Lys	Leu	Leu	Phe	Ile	Ser	Glu	Pro	Ile
				405				:	410					415	-
Pro	His	Pro	Ser	Asn	Glu	Leu	Arg	Gly	Leu	Asn	Glu	Lys	Met	Ser	Asn
			420					425					430		
Lys	Cys	Ser	Met	Leu	Ser	Thr	Ala	Glu	Asp	Asp	Ile	Arg	G1n	Asn	Phe
		435					440					445			
Thr	Gln	Leu	Pro	Leu	His	Lys	Asn	Lys	G1n	Glu	Cys	Ile	Leu	Asp	Ile
	450					455				-	460				
Ser	Glu	His	Thr	Leu	Ser	Glu	Asn	Asp	Leu	Glu	Glu	Leu	Arg	Val	Asp
465					470					475					480
His	Tyr	Lys	Cys	Asn	Ile	Gln	Ala	Ser	Val	His	Val	Ser	Asp	Phe	Ser
•				485				•	490					495	
Thr	Asp	Asn	Ser	G1y	Ser	Gln	Pro	Lys	Gln	Lys	Ser	Asp	Thr	Val	Leu
			500					505					510	)	
Phe	Pro	Ala	Lys	Asp	Leu	Lys	Glu	Lys	Asp	Leu	His	Ser	Ile	Phe	Thr
	•	515					520					525	•		
His	Asp	Ser	Gly	Leu	Ile	Thr	Ile	Asn	Ser	Ser	Gln	Glu	His	Leu	Thr
•	530					535	i				540			1	
Val	Gln	Ala	Lys	Ala	Pro	Phe	His	Thr	Pro	Pro	Glu	Glu	Pro	Asn	Glu
5/5	;				550	1				555	I			•	560

## 52/59

Cys	Asp	Phe	Lys	Asn	Met	Asp	Ser	Leu	Pro	Ser	Gly	Lys	Ile	His	Arg
				565.					570					575	
Lys	Val	Lys	Ile	Ile	Leu	Gly	Arg	Asn	Arg	Lys	Glu	Asn	Leu	Glu	Pro
			580					585					590		
Asn	Ala	Glu	Phe	Asp	Lys	Arg	Thr	Glu	Phe	Ile	Thr	G1n	Glu	Glu	Asr
		595					600					605			
Arg	Ile	Cys	Ser	Ser	Pro	Val	Gln	Ser	Leu	Leu	Asp	Leu	Phe	Gln	Thr
	610					615					620				
Ser	Glu	Glu	Lys	Ser	Glu	Phe	Leu	Gly	Phe	Thr	Ser	Tyr	Thr	Glu	Lys
625					630					635					640
Ser	Gly	Ile	Cys	Asn	Val	Leu	Asp	Ile	Trp	Glu	Glu	Glu	Asn	Ser	Asp
				645					650					655	
Asn	Leu	Leu	Thr	Ala	Phe	Phe	Ser	Ser	Pro	Ser	Thr	Ser	Thr	Phe	Thr
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Gly	Phe														

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<211> 176

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<400> 10 ·

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Lys Lys Ser Ser Thr Ser Val Arg Asp Gly Gly Lys Arg Val Gly Ser
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Gly Ala Gln Lys Thr Arg Thr Gly Arg Leu Lys Lys Pro Phe Val Lys

35

40

45

Val Glu Asp Met Ser Gln Leu Tyr Arg Pro Phe Tyr Leu Gln Leu Thr
50 55 60

Asn Met Pro Phe Ile Asn Tyr Ser Ile Gln Lys Pro Cys Ser Pro Phe
65 70 75 80

Asp Val Asp Lys Pro Ser Ser Met Gln Lys Gln Thr Gln Val Lys Leu

85 90 95

Arg Ile Gln Thr Asp Gly Asp Lys Tyr Gly Gly Thr Ser Ile Gln Leu
100 105 110

Gln Leu Lys Glu Lys Lys Lys Gly Tyr Cys Glu Cys Cys Leu Gln
115 120 125

Lys Tyr Glu Asp Leu Glu Thr His Leu Leu Ser Glu Gln His Arg Asn 130 135 140

Phe Ala Gln Ser Asn Gln Tyr Gln Val Val Asp Asp Ile Val Ser Lys

145

150

155

160

Leu Val Phe Asp Phe Val Glu Tyr Glu Lys Asp Thr Pro Lys Lys Lys 165 170 175

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<223> Description of Artificial Sequence:an artificially
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cccaagcttg acattagata ctacattgaa

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⟨210⟩ 12

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<223> Description of Artificial Sequence:an artificially

## synthesized primer sequence

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30

⟨210⟩ 13

<211> 11

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<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: TAT sequence

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Tyr Gly Arg Lys Lys Arg Arg Gln Arg Arg Arg

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5

10

<210> 14

<211> 12

<212> DNA

<213> Artificial Sequence

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<223> Description of Artificial Sequence:an artificially

## synthesized adaptor sequence

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12

<210> 15

<211> 52

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<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:an artificially
synthesized primer sequence

<400> 15

ataagaatgc ggccgctaag aaggagatat acatatgtac ccctacgacg tg

52

<210> 16

<211> 37

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:an artificially

## synthesized primer sequence

<400> 16

ataagaatgc ggccgcttat cacaagctca tatcttt

37

<210> 17

⟨211⟩ 16

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<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:an artificially
synthesized peptide sequence

<400> 17

Met Tyr Pro Tyr Asp Val Pro Asp Tyr Ala Phe Ser Pro Gln Arg Asp

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15

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<211> 29

<212> DNA

<213> Artificial Sequence

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synthesized primer sequence

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cacggatcca tggcatccag cccggccca

29

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<212> DNA

<213> Artificial Sequence

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<223> Description of Artificial Sequence:an artificially
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<213> Artificial Sequence

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<221> MOD\_RES

<222> (9)

<223> PHOSPHORYLATION

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Cys Arg Gly Asn Asp Pro Leu Thr Ser Ser

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5

10

<210> 21

<211> 10

<212> PRT

<213> Artificial Sequence

⟨220⟩

<223> Description of Artificial Sequence:an artificially
synthesized peptide sequence

**<400> 21** 

Cys Arg Gly Asn Asp Pro Leu Thr Ser Ser

1

5